

FIGURE 1

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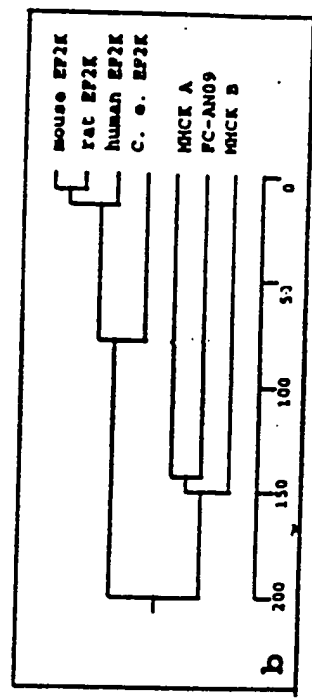
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I
human EP2K 122 SEWLEETLIFMA...
C. o. EP2K 108 KMTST...
MHCK A 570 HWY...
MHCK B 130 A...
PC-AN09 1
consensus

II
human EP2K 178 ---
C. o. EP2K 162 ---
MHCK A 653 ---
MHCK B 177 ---
PC-AN09 42 ---
consensus

III
human EP2K 252 ---
C. o. EP2K 235 ---
MHCK A 734 ---
MHCK B 254 ---
PC-AN09 118 ---
consensus

IV
human EP2K 335 ---
C. o. EP2K 318 ---
MHCK A 811 ---
MHCK B 330 ---
PC-AN09 194 ---
consensus



a

FIGURE 2

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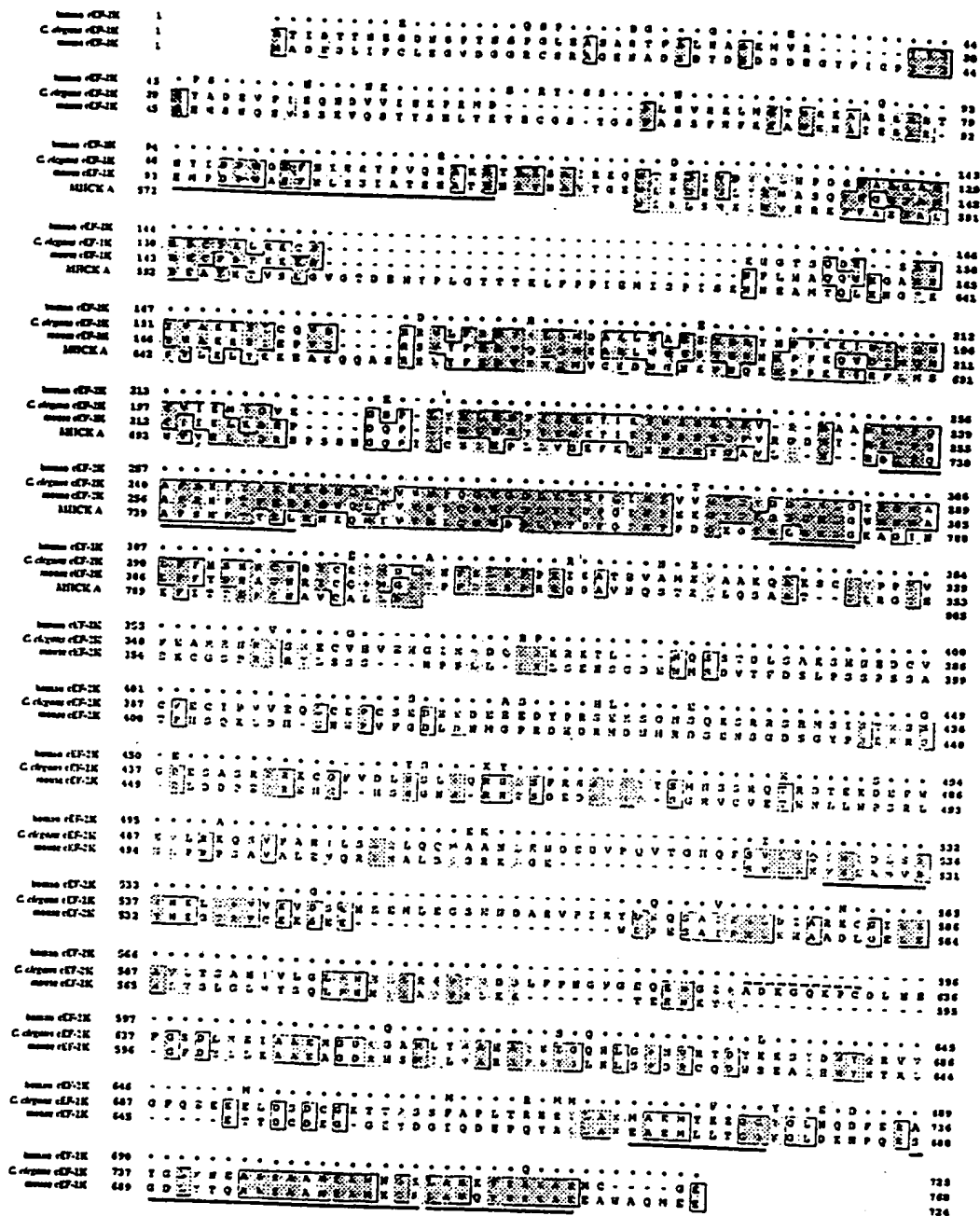
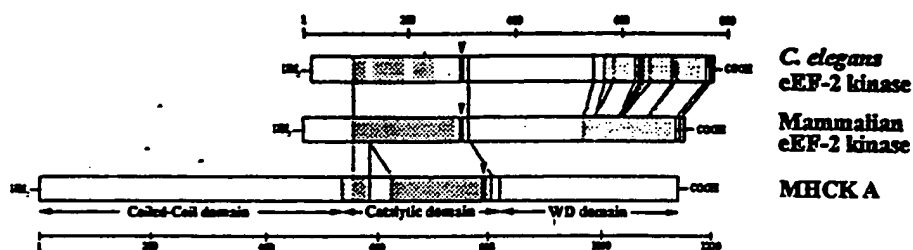


FIGURE 3

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I

H. EF2K 124
C.e. EF2K 110
MHCK A 572
MHCK B 132
MHCK C 48
heart K 189
melano K 48
ch 4 K 1127
consensus

II

H. EF2K 124
C.e. EF2K 110
MHCK A 572
MHCK B 132
MHCK C 48
heart K 189
melano K 48
ch 4 K 1127
consensus

III

H. EF2K 178
C.e. EF2K 162
MHCK A 653
MHCK B 177
MHCK C 89
heart K 249
melano K 108
ch 4 K 1173
consensus

IV

H. EF2K 178
C.e. EF2K 162
MHCK A 653
MHCK B 177
MHCK C 89
heart K 249
melano K 108
ch 4 K 1173
consensus

V

H. EF2K 178
C.e. EF2K 162
MHCK A 653
MHCK B 177
MHCK C 89
heart K 249
melano K 108
ch 4 K 1173
consensus

VI

H. EF2K 252
C.e. EF2K 235
MHCK A 734
MHCK B 254
MHCK C 165
heart K 335
melano K 189
ch 4 K 1253
consensus

VII

H. EF2K 252
C.e. EF2K 235
MHCK A 734
MHCK B 254
MHCK C 165
heart K 335
melano K 189
ch 4 K 1253
consensus

VIII

H. EF2K 252
C.e. EF2K 235
MHCK A 734
MHCK B 254
MHCK C 165
heart K 335
melano K 189
ch 4 K 1253
consensus

FIGURE 5A

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1 cgggcgcggg cgcgtccctc tggccagtca cccggcggag ctggtcgcac aattatgaaa
61 gactcgactt ctgctgctag cgctggagct gagttagttc tgagaagggt tcccggggct
121 gtccctgttc ggtggcccgt gccaccgcct cgggagacgc ttccgatatg gtggctgcag
181 gcgcgggagg tggaggagga gccgtgccc ttccggagtc cgcctcgatga ggagaatgtc
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301 ctccaaagac cctcacagat gtcttccagg atgtcagatt tgtcagcaac ttgtcagatg
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481 gcacacggag cagagcccaa cagatgctta tggagtcac aattttcaag ggggttctca
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6661 aaaattctat aggaatgtgt caatgtgaat tctatttctg gtacttaaga aatcagttgt
6721 tggattatcc ttatacagta tagggagatc acaatacaac tttatgccaa taaaatctaa
6781 cttaattgcc cagatatattt tgcataattt gcaacaagaa aagcttatca tttgactcaa
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7081 tgtttacaaa

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FIGURE 5C

MSQKSWIESTLTKRECVYIIPSSKDPHRCLPGCQICQQLVRCFCGRLVKQHACFTASLAM
KYSDVRLGEHFNQAIIEWSVEKHTEQSPTDAYGVINFQGGSHSYRAKYVRLSYDTKPEII
LQLLLKEWQMELPKLVISVHGGMQKFELHPRIKQLLGKGLIKA AVTTGAWILTGGVNTGV
AKHVGDALKEHASRSSRKICTIGIAPWGVNIENRNDLVGRDVVAPYQTLNPLSKLNVLNN
LHSHFILVDCGTVGKYGAEVRLRRELEKTINQQRIHARIGQGVVVALIFEGGPNVILTV
LEYLQESPPVPVVVCEGTGRAADLLAYIHKQTEEGGNLPDAAEPDIISTIKTFNFGQSE
AVHLFQTMMECMKKKELITVFHIGSEDHQDIDVAILTALLKGTNASAFDQLILTAWDRV
DIAKNHV FVYGQOWL VGSLEQAMLDALVMDRVSVFKLLIENGVS MHKFLTIPRLEELYNT
KQGPTNPMLFHLIRDVKQGNLPPGYKITLIDIGLVIEYLMGGTYRCTYTRKRFRLIYNSL
GGNNRRSGRNTSSSTPQLRKSHETFGNRADKKEKMRHNNFIKTAQPYRPKMDASMEEGKK
KRTKDEIVDIDDPEIKRFPYPLNELLIWACL MKRQVMARFLWQHGEESMAKALVACKIYR
SMAYEAKQSDLVDDTSEELKQYSNDFGQLAVELLEQSFRQDETAMKLLTYELKNWSNST
CLKLAVSSRLRPFVAHTCTQMLLSDMWMGRLNMRKNSWYKVILSILVPPAILMLEYKTKA
EMSHIPQSQDAHQMTMEDSEN NFHNITEEIPMEVFKEVKILDSSDGKNEMEIHKSKKL P
ITRKFYAFYHAPIVKFWFNTLAYLGFLMLYTFVVLVKMEQLPSVQEWIVIAIYIFTYAIEK
VREVF MSEAGKISQKIKVWFSDFNVSDTIAIISFFVGFLRFGAKWNYINAYDNHVFVA
GRLIYCLNII FVYVRLLD FLAVNQAGPYVMMIGK MVANMFYIVVIMALVLLSFGVPRKA
ILYPHEEPSWSLAKDIVFHPYWMIFGEVYAYEIDVCANDSTLPTICGPGTWLTPFLQAVY
LFVQYIIMVNLLIAFFNNVYLQVKAISNIVWKYQRYHFIMAYHEKPVLPPLIILSHIVS
LFCCVCKRRKKDKTSDGPKLFLTEEDQKKLHDFEEQCVEMYFDEKDDKFNSGSEERIRVT
FERVEQMSIQIKEVGDRVNYIKRSLQSLDSQIGHLQDLSALTVDTLKTLTAQKASEASKV
HNEITRELSISKHLAQNLIDDVPVRPLWKKPSAVNTLSSSLPQGDRESNNPFLCNIFMKD
EKDPQYNLFGQDL PVI PQRKEFNIPEAGSSCGALFPSAVSPPELRQRRHGVEMLKIFNKN
QKLGS SPNSSPHMSSPPTKFSVSTPSQPSCKSHLESTTKDQEPIFYKAAEGDNIEFGAFV
GHRDSMDLQRFKETS NKIRELLSNDTPENTLKHVGAAGYSECKTSTSLHSVQAESCRR

FIGURE 6B

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ASTEDSPEVDSKAALLPDWLRDRPSNREMPSEGGTLNGLASPFKPVLDNTNYYSAVERN
LMRLSQSIPFVPPRGEPTVYRLEESSPSILNNSMSSWSQLGLCAKIEFLSKEEMGGG
LRRVAVKVLCTWSEHDILKSGHLYI IKSFLPEVINTWSSIIYKEDTVLHCLREIQQRAAQ
KLTFAFNQMKPKSIPYSPRFLEVFLLYCHSAGQWFAVEECMTGEFRKYNNNGDEI IPTN
TLEEIMLAFSHWTYETRGELLVLDLQGVGENLTDPSVIKAEERSCDMVFGPANLGEDA
IKNFRAKHHHCNSCCRKLKLPDLKRNDYTPDKIIFPQDESSDLNLQSGNSTKESEATNSVR
LML

Figure 7A

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[illegible]

MSQKSWIESTLTKRECVYIIPSSKDPHRCLPGCQICQQLVRCFCGRLVKQHACFTASLAM
KYSDVKLGDHFNQAIEEWSVEKHTEQSPTDAYGVINFQGGSHSYRAKYVRLSYDTKPEV
ILQLLLKEWQMELPKLVISVHGGMQKFELHPRIKQLLGKGLIKAAVTTGAWILTGGVNT
GVAKHVGDALKEHASRSSRKICTIGIAPWGVNIENRNDLVGRDVVAPYQTLNPLSKLNV
LNNLHSHFILVDDGTVGKYGAEVRLRRELEKTINQQRIHARIGQGVVVALIFEGGPNVIL
TVLEYLQESPPVPVVCEGTGRAADLLAYHKQTEEGGNLPDAAEPDIISTIKKTFNFGQN
EALHLFQTLMECMKRKELITVFHIGSDEHQDIDVAILTALLKGTNASAFDQLILTLAWDR
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NSLGGNNRRSGRNTSSSTPQLRKSHESFGNRADKKEKMRHNFHFIKTAQPYRPKIDTVME
EGKKKRTKDEIVDIDDPETKRFPYPLNELLIWACLMKRQVMARFLWQHGEESMAKALV
ACKIYRSMAYEAKQSDLVDDTSEELKQYSNDFGQLAVELLEQSFRQDETMMAMKLLTYE
LKNWSNSTCLKLAVAAKHRDFIAHTCSQMLLTDMWMGRLMRKNPGLKVILSILVPPAI
LLEKYKTKAEMSHIPQSQDAHQMTMDDSENNFQNITEEIPMEVFKEVRILDSNEGKNEM
EIQMKSKKLPIRTRKFYAFYHAPIVKFWFNTLAYLGFLMLYTFVVLVQMEQLPSVQEWIVI
AYIFTYAIEKVREIFMSEAGKVNQKIKVWFSDYFNISDTIAIISFFIGFLRFGAKWNFANA
YDNHVFVAGRLIYCLNIIFWYVRLLDFLAVNQQAGPYVMMIGKMMANMFYIVVIMALV
LLSFGVPRKAILYPHEAPSWTLAKDIVFHPYWMIFGEVYAYEIDVCANDSVIPQICGPGT
WLTPLQAVYLFVQYIIMVNLLIAFFNNVYLQVKAISNIVWKYQRYHFIMAYHEKPVLP
PLIILSHIVSLFCCICKRRKKDKTSDGPKLFLTEEDQKKLHDFEEQCVEMYFNEKDDKFHS
GSEERIRVTFERVEQMCIQIKEVGDRVNYIKRSLQSLDSQIGHLQDLSALTVDTLTKTLTAQ
KASEASKVHNEITRELSISKHLAQNLIDDGPVRPSVWKKHGVVNTLSSSLPQGDLESNNP
FHCNILMKDDKDPQCNIFGQDLPAVPQRKEFNPEAGSSSGALFPSAVSPPELRQRLHGV
ELLKIFNKNQKLGSSTSIPLHSSPPTKFFVSTPSQPSCKSHLETGTKDQETVCSKATEGDN
TEFGAFVGHRSMDLQRFKETSNIKILSNNNTSENTLKRVSLSLAGFTDCHRTSIPVHSKQ
EKISRRPSTEDTHEVDSKAALIPVWLQDRPSNREMPSEEGTLNGLTSPFKPAMDTNYYYSS
AVERNLMRLSQSIPFTPVPPRGEPVTYRLEESSPNILNNSMSSWSQLGLCAKIEFLSKE
EMGGGLRRAVKVQCTWSEHDILKSGHLYIISFLPEVVNTWSSIYKEDTVLHLCLREIQQ
QRAAQKLTFAFNQMKPKSIPYSPRFLEVFLLYCHSAGQWFAVEECMTGEF
RKYNNNNGDEIPTNTLEEIMLAFSHWTYETRGELLVLDLQGVGENLTDPSVIKAEKR
SCDMVFGPANLGEDAIGNFRAKHHCNSCCRKLKLPDLKRNDYTPDKIIFPQDEPSDLNLQ
PGNSTKESESTNSVRLML

Figure 7B

[illegible]

Figure 8A

ESAEPLTQSDKRETSHTTAAATGRSSHADARECAISTAQAEQAKTLQTSTDSVSKEGNTNCKGEGMQVN
TLFETSQVPDWSDPPQVQVQETVRETISCSQMPAFSEPAAGEESPFTGTTISFNLGGVHKENASLAQHSEV
KPCTCGPQEEKQDRDGNIPDNFREDLKYEQISEANDETMSPGVFSRHLPKDARADFREPVAVSVASP
TDTALTLENVCDEPRDREAVCAMECFEASDQGTCTFDITDSL VGTPVDNYSQEQICSVDTTELAEQONKVS
LCSSNDKTLVFFQTQVSETSVTCKSSKDGNSVMSPLFISTFTLNISHTASEGATGENLAKVEKSTYPLAS
TVHAGQEPPSPNSGGLDETQLLSENPNLVQFKEGGDKSPSAADTTATPASYSSIVSFPWEKPTTLTAN
NECFQATRETVTIATEVHPAKYLA VSIPEDKHAGGTEERFPRA SHEKVSQFPSQVQVDHILSGATIKSTKEL
LCRAPSVPGVPHHVLQLPEGEFCNSPLQVDNLSGDKSQTVDRADFRSYEENFQERGETKQGVQQSL
SQQGLSAPDFQQSLPTTSA AQEERNLVPTAPSPASSREGAGQRSGWGTRVSVVAETA GEEDSQALSNVPS
LSDILLEESKEYRPGNWEAGNKLKIITLEASASEIWP RPRLTNSESKASDGGIIPDKVWAVPDSLKADAVV
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AASETGGKENVNNVSQDQEEKQLKMDHTAFFKKFLTCPKILESSVDPIDEISVIEYTRAGKPEPSETTPQGA
REGGQSNNDGNMGHEAEIQSAILQVPCLOQTILSENRISSRQEGSMKQEAEQIPEEAKTAIWQVLQPSEGG
ERIPSGCSIGQIQESSDGLGEAEQSKKDKAELISPTSPSSCLPIMTHSSLGVDTHNSTGQIHDVPENDIVER
RKRQYVFPVSQKRGTIENERGKPLPSSPDLTRFPCTSSPEGNVTDFLISHKMEEPKIEVLQIGETKPPSSSSS
AKTLAFISGERELEKAPKLLQDPCQKGTLCACKSREREKSLEARAGKSPGTLTAVTGSEEVKRRKPEAP
GHLAEGVKKKILSRVAALRLKLEEKENIRKNSAFLKMPKLETSLSHTEEKQDPKPKPSCKREGRAPVLLK
KIQAE MFPEHSGNVKLS CQFAEIHEDSTICWTKDSKSI AQVQRSAGDNSTVSFAIVQASPKDQGLYYCCIK
NSYGKVTAEFNLTAEVLKQLSSRQDTKGCEEIEFSQLIFKEDFLHDSYFGGRLRGQIA TEELHFEGEVHRK
AFRSTVMHGLMPVFKPGHACVLKVHNAIAYGTRNNDDELQIRNYKLAQECYVQNTARYYAKIYAAEAQ
PLEGFGEVPEIPIFLIHRPENNIPIYATVEEELIGEFVKYSIRDGKEINFLRESEAGQKCCCTFQHWVYQKTS
CLLVTDMMQGVGMKLT DVGIA TLAKGYKGFKGNCMSMTFIDQFKALHQC NKYCKMLGLKSLQNNNQKQK
QPSIGKSKVQTN SMTVKKAGPETPGEKKT

Figure 8B

1 atgtccacaga aatcctggat taaaggagta ttgacaaga gagaatgtag cacaatcata
 61 cccagctcaa aaaatcctca cagatgtact ccagtatgcc aagtctgccaa gaatttaalc
 121 aggtgttact gtggccgact gattggagac catgtctgga tagattatc ctggaccalc
 181 tcagctgccaa agggtaaaga aagtgaacaa tggctgttg aaaagcacac aacgaaaagc
 241 ccaacagata cttttggac gattaattc caagatggag agcacacca tcatgccaa
 301 tataatagaa cttctatga taaaaactg gatcaictgt tacatttaac gtgaaagag
 361 tggaaaatgg aactgcccac gctgtgac tcagtccatg ggggcalca gaactttact
 421 atgcccctca aatttaaga gattttcagc caaggtttg taaagctgc agagacaaca
 481 ggagcgtgga taataactga aggcacat aacagtgtcca agcatgttg ggaatgcttg
 541 aaatcccat cctctcattc ctgagaaaa atctggacag ttggaatccc tcttggggt
 601 gtcaatgaga accagagaga cttatggga aaagatgtg tgtccctga ccagactctg
 661 gataaccccc tcagcaagct cacaacact aacagcatgc actgcactt catcctgtct
 721 gatgatggga cgttgggcaa gtatggaat gaaatgaagc tcagaaggaa cctggagaag
 781 taccctctc tgcagaaaa acactgccgc tcaagacaag gctgcccgt cgtgggctg
 841 gtgttggag gcgggtccaa cgtatcctg tcagtgtgg agactgtcaa ggacaaggac
 901 ccagtgttg tgttgaggg cacaggtagg gcggctgacc tctggcctt cacacacaaa
 961 cacttgccag atgaaggat gctgcgacct caggtgaag aggaatcat ctgcatgatt
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 1441 catctctgc aagatgtga acagcalacc ctttttcag gctaccgaat aaccttgatt
 1501 gacattgat tagtagtaga atacctcatt gtagagcat atgcagcaa ctacactaga
 1561 aaacattica gagccctcta caacaacctc tacagaaaa acaagcacca gagacactcc
 1621 tcaggaaaata gaaatgagtc tgcagaaagt acgtctgact cccagttcat tagaactgca
 1681 cagccataca aattcaagga aaagtctata gtccctcata aatcaaggaa gaagcaaaa
 1741 gaacaaaatg tatcagatga cctgagctc actggcttc ttaccctta caatgacctg
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 2581 gagtggctg ttacattta catctcacc aatgtattg aggtggcag ggaggtgagt
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 2701 ttaacagaaa ctgtggccat tggcctgttt tcagctgct tctctcttg atgggtgac
 2761 cctcttttc acacagcggg aagactgac tactgcatag acatcatatt ctggttcca
 2821 cggctcttg acttttgc tgtaatcaa catgcaggtc cataatgac catgatgca
 2881 aaaatgacag caaacatgt ctatattg atcatatgg ccatagtct cgtgagctt
 2941 ggagtgacac gcaaggccat cctttgcca aaagagccac catctggag tctagctga
 3001 gatattgtat ttgagccata ctggatgata tacggagaag tctatgttg agaaatagat
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 3121 gttactctc tctgcaata taltcatg gtgaacctgt tgattgctt cttaacaa
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 3241 atcatgacct accagagaa gccctggctg cccccacct tcatctgtc gagccacgtg
 3301 ggcttttcc tccgcccct gtgtgtcat cagctcctc acgaccaaga agagggtgac
 3361 gtggattaa aactctacct cagtaaggag gatctgaaa aacttcatga tttgaggag
 3421 cagtgcgtg aaaaatact ccatgagaag atggaagatg tgaattgag ttgtaggaa
 3481 cgaatccgag tgacatcaga aagggttaca gagaatgact tccagctgaa agaaatgaa
 3541 gaaaaggtgt ctttataaa ggaactctta ctgtctttg acagccaggt gggacacctg
 3601 caggatctct ctgcccctgac tgtgatacc ctgaaagtc ttctgtctg tgacacttg
 3661 caagaggatg aggtctctc ggccaagaga aagcatttca ctgcaaaa acttccccac
 3721 agctggagca atgtcaictg tgcagaggt ctaggcagca tggagatgc tggagagaag
 3781 aaataccagt attatagcat gccctctct tgcagaga gccctggctg aggccggcat
 3841 cccccaagag tgcagagggg ggcacttct gagattacaa acagtaaaag agaggctaca
 3901 aatgtaagaa atgaccagga aaggcaagaa acacaagta gtatagtgt ttctggggtg
 3961 tctctaaca ggcaagcaca ctcaaatgt ggccagttc ttctgtccc ctctaacta

Figure 9A

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4021 aagcgagttc cttttcagc agaaactgtc ttgcctctgt ccagaccctc tgtgccagat
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4141 ccagtgtct ctgactgggc aicagtggat gaaccaagc aaaagcacga gctatttct
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4261 gaacccaatga caatgagctc cctctttcc caagccaaga tcatgcaaac tggagggtga
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4681 aaagcgaaaa tgcataacaa agacaggaga ctgtcaaga aaaagaaga tctcaagga
4741 tccagggtgc caatcaaac agtcaagcc tgcctcaga gtgaccagt gaalccagag
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5221 gataaaagca tgcctcttg gtctcagcgt gggagagcgg caatgatcca ggtattgtcc
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5341 gatgacattc tcaagccggg acaagtttc altgtcaagt cctttctcc tgggtgtg
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5521 atacctata caccaaggtt cctggaagt ttctaatct actgcattc agccaaccag
5581 tggttgacca ttgagaagta tatgacagg gagttccgga agtatacaa caacattgt
5641 gatgaaatca cccccacaa caccctggag gagctgatgt tggcttctc tcactggacc
5701 taigagtaca ctgggggaga gctgtgtgt ttgatttgc aaggtgttg agaaaattg
5761 acagatccat ctgttataa acctgaaagc aaacaatcaa gaggaatgt gttggaccg
5821 gccaatitgg gggaaagatc aattagaac ttcatgcaa aacatattg taactctgc
5881 tggcgggaagc tcaaatctcc ggaattaaa agaaatgact attccctga aagataaat
5941 tccacctttg gacttgagat aaaaatagaa tcagctgagg agcctccagc aaggagagc
6001 ggtagaaatt cccagaaaga tgaatgcaa ctataaaaag gggaggacaa gaagatccca
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6241 agctccatcc tcaagtcca caggaagatg ctggatgag tcaagctgga atattgtct
6301 tgtgtacct attgcttag ctgtcactt ggaacttgg agcagaatcc tgcattaa
6361 aggatgggg tgggggggat acattattt tatttttca ctatgtatc agactggacc
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6481 aaaagttagt tgattctc ctctgtctt tccacaacat aggaattga atagcaatga
6541 taggaaaaac aatggaacaa ggggtgggtt gcacagattg gagcacattc ctgcacaaac
6601 taccagatct actggtgaaa tctgatggg ttccagatat tgcagtgaa tcatatgat
6661 ctggatattt tcaagtttct gtaaaagaaa gggaaacctt aaacaatac ccttcatal
6721 ataataata tggaaatgt atatatata tattttata tatataat atattgaata
6781 tatataatt atatatataa tacaatgga atatatata ttatatata tatatatat
6841 ttattttg agatggagt tcaactttt taccaggct ggagtgcatt gatcgatct
6901 cactgcaacc tctgcctccc gggcttgagc gattcttgc tctcagctt cgggttagct
6961 gggactacag gtgtgacca ctatgcctgg ctattttgt atttttagt gagatgggt
7021 ttaccatgt tggccaggct ggttcaaac tctgacctc agatgatcca cctgccttg
7081 ctcccaag tctgggatt acaggcgtga gccactgcgc ctggccttt ttttttt
7141 tttaacgag aacaagaata tgaagaactg gaaatcatta agaaagggt tccctctct
7201 aaagctcagg ggtactatta gttaggagt gactaactca acctgtaaa caccactct
7261 ccttccaaag ttgtatatat aatattgcag gttaaattac ttatgtcag gtctatgaa
7321 gaaagatacg gtttcagact gaaaacatgt ttccagggt ttgtctct tccagagcag
7381 agttccctat tccctggga taaagaatgt atatatatt tgaatatgg ctgagaacat
7441 gatttggtt tgtaggcct aagtggaagc actcctggca gccacactgt gtatgtatt
7501 tgagggatca gtatccctc ttgtatctg ggcctgtgt cctacactg aacaagcacc
7561 agctttcac acaaggagag atgtggggct gggagtctc tccccactc attgatctc
7621 cttcttatt ataagctgt ccagttcaca ggcagcaaac ctctgggtt tgaataatc
7681 caactattt ttatcttaa tctgacatt agctgactg ctagtgtct tctttaaa
7741 atctacactc ttgactctt aggcatacag gggaaatgt gaaaaggag gtggaaaacc
7801 aagaatttag ttgccaatg attgcctctg attctgtaa gtttgagtc cacaaggct
7861 aatttatcc cctttactt ggggtttgg gtgtggaaa gcgggaatt tgggtgatt
7921 gttgatggc aatgaggata aaatgtaat actttttgg ggaattaca actttatct
7981 attctacaag tcaataagg acaattgtt actcacctca gtgtgcact caatatgga

Figure 9A

601-1-0984P

8041 aagaggcaga gttgtctgc ccaattgeca aactaaagac atcagttcat tggtaata
8101 ttgttacct ggaatggaac ttgaagcaa atacattgg atticaaatt tcaaaaaa

Figure 9A

[illegible]

Human kidney kinase

MSQKSWIKGVFDKRECSTIIPSSKNPHRCTPVCQVCQNLIRCYCGRLIGDHAGIDYSWTIS
 AAKGKESEQWSVEKHTTKSPTDTFGTINFQDGEHTHHAKYIRTSYDTKLDHLLHMLKE
 WKMELPKLVISVHGGIQNFTMPSKFKEIFSQGLVKA AETTGAWIITEGINTVSKHVGDAL
 KSHSSHSLRKIWTVGIPPWGVNIENQRDLIGKDVVCLYQTLNPLSKLTTLNSMHSFILS
 DDGTVGKYGNEMKLRRLNLEKYL SLQKIHCRSRQGVVGLVVEGGPNVILSVWETVKD
 KDPVVVCEGTGRAADLLAFTHKHLADEGMLRPQVKEEIIICMIQNTFNFSLKQSKHLFQIL
 MECMVHRDCITIFDADSEEQQDLDLAILTALLKGTNLSASEQLNLAMAWDRVDIAKKHI
 LIYEQHWKPDALQAMSDALVMDRVDFVKLLIEYGVNLHRFLTIPRLEELYN TKQGPTN
 TLLHHLVQDVKQHTLLSGYRITLIDIGLVVEYLIGRAYRSNYTRKHFRALYNNLYRKYK
 HQRHSSGNRNESAESTLHSQFIRTAQPYKFKEKSIVLHKSRRKSKEQNVSDDPESTGFLY
 PYNDLLVWAVLMKRQKMAMFFWQHGEETVKA VIACILYRAMAHEAKESHMVDDAS
 EELKNYSKQFGQLALDLEKAFKQNERMAMTLLTYELRNWSNSTCLKLAVSGGLRPV
 SHTCTQMLLTDMWWMGR LKMRKNSWLKIIISILPPTILTLEFKSKAEMSHVPQSQDFQFM
 WYYSDQNASSSKESASVKEYDLERGHDEKLDENQHFGLSEGHQHLPWTRKVYEFYSAP
 IVKFWFYTMAYLAFLMLFTYTVL VEMQPQPSVQEWLVSIYIFTNAIEVVREVSISEPGKF
 TQKV KWISEYWNLTETVAIGLFSAGFVLRWGDPPFHTAGR LIYCIDIIWFWSRLLDFFA
 VNQHAGPYVTMI AKMTANMFYTVIIMAI VLLSFGVARKAILSPKEPPSWSLARDIVFEPY
 WMIYGEVYAGEIDVCSSQPSCPPGSFLT PFLQAVYLFVQYIIMVNLLIAFFNNVYLDMESI
 SNNLWKYNRYRYIMTYHEKPWLPPPLILLSHVGLLLRRLCCHRAPHDQEEGDVGLKLY
 LSKEDLKKLHDFEEQCVEKYFHEKMEDVNCSCEERIRVTSE RVTEMYFQLKEMNEKVS
 FIKDSL LSLDSQVGH LQDLSALTVDTLKVL SAVDTLQEDEALLAKRKHSTCKKLPHSWS
 NVICAEVLGSMEIAGEKKYQYYSMPSSLLRSLAGGRHPPRVQRGALLEITNSKREATNV
 RNDQERQETQSSIVVSGVSPNRQAHSKYGQFLLVPSNLKRVPFSAETVLPLSRPSVPDVL
 ATEQDIQTEVLVHLTGQTPVVSDWASVDEPKEKHEPIAHL LDGQDKAEQVLPTLSCTPE
 PMTMSSPLSQAKIMQTGGGYVNWAFSEGDETG VFSIKKKWQTCLPSTCDS DSSRSEQHQ
 KQAQDSSLSDNSTRSAQSSECSEVGPWLQPN TSFWINPLRRYRPFARSHSFRFHKEEKL
 KICKIKNLSGSSEIGQGA WVKAKMLTKDRRLSKKKKNTQGLQVPIITVNACSQSDQLNP
 EPGENSISEEEYSKNWFTVSKFSHTGVEPYI HQKMKTK EIGQCAIQISDY LKQSQEDLSKN
 SLWNSRSTNLNRNSLLKSSIGVDKISASLKS PQEPHHHYS AIERNNLMRLSQTIPFTPVQL
 FAGEEITVYRLEESSPLNLDKSMSSWSQRGRAAMIQVLSREEMDGG LRKAMRVVSTWS
 EDDILKPGQVFIVKSF LPEVVRTWHKIFQESTVLHLCLREIQQQRAAQKLIYTFNQVKPQ
 TIPYTPRFLEVFLIYCHSANQWL TIEKYMTGEFRKYNNNNGDEITPTNTLEELMLAFSHW
 TYEYTRGELLVLDLQGVGENLTDPSVIKPEVKQSRGMVFGPANLGEDAIRNFI AKHHCN
 SCCRKLKLPDLKRNDYSPERINSTFGLEIKIESAEPPARETGRNSPEDDMQL

Figure 9B

[illegible]

Figure 10A

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CTTGTACCCCTCGCCACATCCCTGTCTTGGGGCCAGCTGCAGCTGCTGCTGCCCTCTGAGTGTTCACGCTCTGACATGGCCCTTGTCTGAGGGTCCCGACCTCCCTCG
CCACCAATACCCAGGTGAGGAACAGACCTCTGGCTCTCACCCCACTTCAAGTCTCTTCCCAACTTCTCTCGGGCTCTTTGCTCATGAGGTGAGAGCTGGTG
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CTCTGGGAATTAGGACCCCTAGTCCCAACCATCGCTCTGATCTTGGGGCCCGGCTCTGGTCTCTCATGATGGCTCCCAAGGCCAGCCCTGGATCTCTTC
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MEVAWL VYVLGQQPLARQGEQSRQLVPGRGLVLWLPGLPRSSPSWPAVDLAPARPRGLICHTGHEQAGREPQGSST
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RGARQPRAGAAAAGRPGAGAGWRTGEAAAASAGPAV GEGGAMGSRRAPTRGWAGGRSGAGDGEDDGPVWIPSPASRS
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DGEHGLLTYICDAMELGPQRALKEESGAKKKKDEESKQGLRKPELEKAAQSRSSENCIPSSDEPDSCGTQGPVGVQVQT
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APLRARSEGVPGAPGQPTHSLTPQPTRPFNRKRFAPPKPKGEATTDSPISLSQAPECGAQSGLKAPQASVQVPTPPARRRH
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QESSMAGRLGEAGGQAAPGQGPSAESIAQEPSQEEKFPGEALTGLPAAATPEELALGARRKRFLPKVRAAGDGEATTPEERES
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LKAPQVIRKIRVEQFPDASGSLKLWCQFFNILSDSVLTWAKDQRPVGEVGRSAGDEGPAAALIVQASPVDCGVYRCTIHNEH
GSASTDCLSPVLSGFISREEGEVGEIEMTMPVFAKGLADSGCWGDKLFGRLVSEELRGGGYGCGLRKASQAKVIYGLEPI
FESGRTCIHKVSSLLVFGPSSSETSLVGRNYDVTIQCKIQNMSREYCKIFAAEARAAPGFGEVPEIIPLYLIYRPANNIPYATLEE
DLGKPLESYCSREWGC AEAPTASGSSEAMQKCCQTFQHWLYQWWTNGSFLVTDLAGVDWKMTDVQIATKL RGYQGLKESCF
PALLDRAFSSHQCNAYCELLGLTPLKGP EAAHPQAKAKGSKSPSAGRKGSQ LSPQPKKGLPS PQGTRKSA PSSKATPQASEP
VTTQLLGPPTQEEGSKAQGM R

Figure 10B

ATGAATAATCAAAAAGTGGTAGCTGTACTGCAAGAGTGCAAGCAAGTGTGGATCAGCTCTTGTGGAAAGCGCCAGATGTGTGGAAAGAGGACAAGAGCGGA
GGACCAGCGCTGCAGAGCTTTACTTCCCGCAGGAGTTAAGGACCTGTATCCAGGAGGCAAGGAAATGAAGTGGCCCTTCGTGCTCGCTGAAAAGTGGCAGTACAAAC
AAGCCGTGGGCCCGCAGAGGACAAACAACCTGAAGGATGTGATTGGCCCGGGTTGCAAGAGTTACTGGCGTCCCTGAGGGCTCCATCTCGCTCGCGGACTGT
CCAGCCCAATTGCCCGCAGGTGTTATTAGCCCAAGCCCGAATCTCCGTGAACCTCAGGAATACTTTAAAGCAGAGTATATCTGAGCAGTCTAATAAGCAAC
ATGGAGCAACGGGTACCTGGCTGTACAGAAATGAAGGAAATCTCGGTGAACCTCAGAGTCTGTAACAGATCAGAGGGCAGATTCTGCAAAAAGCTGGGTAT
TGGTACGAAGCAGCAGAGTTAATATGGGCCCTCCATGTAGGATATTGGCAGATTCTCCAGCCGGAATAAAGGGCTCTCCAGCTCGCTAGGTATATCTGGCAG
ATCTTTGTTTCCAAGAGCAAGCAAGATTATGAAGAGTTTAAAGAAATCCACAATTAATGTAGCGCTGTGAAGGAGTTTGACCACTTATGCTTCCGCTCGCAG
AAGCTTCAAGCTGGCAGCTGCCTCAGTGGCTATACGCCGCTCTTCGTCTCACAGCTGTGAATCCGTGGCCACGTGTATGTGCTCTACAGTGTCTCAAAATGA
GTCTCTCCAGATTGAAAACCTTACATCTGTGTGAAGCCAAAGAGGCCCTTGAGATGGCTCTCCACCAAGAGAGATGATGAGCCCTGTATCTGCTACAGTGTCTCAAAATGA
CTTTCACAGATTGTCAAACTTTCGGTCTCCACAGTGCACAGAAAGGCTCCATGGGGAGCAGGAGCGTCCATCGACGAAGTATATGTCTGTGATGGCCAGGTGAAGGAAGC
AATGGGAAGCTGTACAAATTCAGCACTTCTCCAGAGTCAGCAAGAGAGAGCTGTCTGTCAAGAAAGTATATGTCTGTGATGGCCAGGTGAAGGAACATTTTACA
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AAGATGTTCAGAGGAACTCAGAAAGGGGAGGAGGAGAACTGGACCCCATCTGATGCAATTCGAGTCTCTTGGATCAAGATGTGGAGACTGAGACTGAGCC
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AGGAAGTCAGAAATAGGGACCCAGAAATACTTGTGCTCAGACCCCTCATATGTTCTGTCTTGGTCTCTGATCTGGTAGGGCCCAAGAAATATGGGCAC
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TGCTTTTGGGTTCATCATCTCAAGAAAGAAATCTGGGAGGTATGTGGGAAGACTATAAGGAGCAGAGGGGCTCTGGCACCTTCACTGATGTGGA
CGCAGATGACCGCAGCACTATGTGACAGAAATTAACAAGAGACTATGAACAAAAACATCCCAACCCAGATATCTACATCCCATCCCAATACTACTGAT
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GAATGTAATGAAATCTGCCATCGTCTTTTGACTAGACCTTCAATGGGAGAAACCA

Figure 11A

MNNQKVAVLLQECKQVLDQLLEAPDVSEEDKSEDQRCRALLPSELRTLQEAKEMKWPFPVEKWQY
 KQAVGPEDKTNLKDVGAGLQQLLASLRASILARDCAAAAIVFLVDRFLYGLDVSGKLLQVAKGLHKL
 QPATPIAPQVVIRQARISVNSGKLLKAEYILSSLSNNGATGTWLYRNESDKVLVQSVCIQIRGQILQKLG
 WYEAELIWAIVGYLALPQPDKKGLSTSLGILADIFVSMKNDYEKFKNPQINLSLLKEFDHHLSSAA
 ACKLAAAFSAYTPLFVLTAVNIRGTCLLSYSSSNDPPELKNLHLCEAKEAFEIGLLTKRDDDEPVTGKQEL
 HSFVKAAGFLTTVHRRLLHGETGTVHAASQLCKEAMGKLYNFSTSSRSQDREALSQEVMSVIAQVKEHLQ
 VQSFNSVDDRSYVPESFECRLDKLILHGQGFQKILDY'SQHHTSVCEVFESDCGNNKNEQKDAKTGVCI
 TALKTEIKNIDTVSTTQEKPHCQRTDGISSSLMGKNVQRELRRGGRNWTHTSDAFRVSLDQDQVETETEPSD
 YSNGEGA VFNKSLSGSQTSASWSNLGFFSSASWEEVNYHVDDRSARKEPGEHLVDTQCSTALSEELEN
 DREGAMHSLHSLHDLSLQEPNNDNLEPSQNQQQMPPLTPFSPHNTPGIFLAPGAGLLEGAPEGIQEVR
 NMGPRNTSAHSRPSYRSASWSSDSGRPKNMGTHPSVQKEEAFEIIVEFPETNCDVKDRQKEQGEEISERG
 AGPTFKASPSWVDPEGETAESTEDAPLDFHRVLHNSLGNISMLPCSSFTPNWPVQNPDSRKSGGPVAEQGI
 DPDASTVDEEGQLLDSMDVPCNTNGHSHRLCILRQPPGQRAETPNSSVSGNILFPVLSRCTTTEEGNQPG
 NMLNCSQNSSSSVWWLKSPAFSSGSSEGDSPWSYLNSSGSSWVSLPGKMRKEILEARTLQPDDEFKLLA
 GVRHDWLFQRLNTGVFKPSQLHRAHSALLKYSKKSELWTAQETIVYLG DYLT VKKKGRQRNFAFWH
 HLHQEEILGRYVVGKDYKEQKGLWHHFTDVERQMTAQHYVTEFNKRLYEQNIPTQIFYIPSTILLEEDKTIK
 GCISVEPYILGEFVKLSNNTKVVKTEYKATEYGLAYGHFSYEFNSNHRDVVVDLQGWVTGNGKGLIYLT
 QIHSVDQKVFTTNFGKRGIFYFFNNQHVEECNEICHRLSLTRPSMEKP

Figure 11B

HeEF-2_kinase> 1 LDD--EQLDCAQPPFGR-----EFCSTTKGSSNF--WKGASN--AARIEPDR--DVEF
 MHCK_B> 1 ECT--ATLQRPVPFAR-----EFTLTDES-----KSGASGR--EIGKPTPR--PSFP
 Melanoma_kinase> 1 EQLGLCAKQFLDREMG-----EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Kidney_kinase> 1 EORGRAAGQFLDREMG-----EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Muscle_kinase> 1 EOK--EFGRIVEELRCGYCC--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Heart_kinase> 1 EOK--EFGRIVEELRCGYCC--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Lymphocyte_kinase> 1 EOK--EFGRIVEELRCGYCC--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 consensus 1 EOK--EFGRIVEELRCGYCC--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 1 vt vimse G amRkafr l i g yvik y v y ed l

HeEF-2_kinase> 62 MEILGERTRRHPKQ--VHTMCCTIDEKD--P-GKPLEH--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 MHCK_B> 56 MEILGERTRRHPKQ--VHTMCCTIDEKD--P-GKPLEH--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Melanoma_kinase> 74 MEILGERTRRHPKQ--VHTMCCTIDEKD--P-GKPLEH--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Kidney_kinase> 74 MEILGERTRRHPKQ--VHTMCCTIDEKD--P-GKPLEH--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Muscle_kinase> 75 MEILGERTRRHPKQ--VHTMCCTIDEKD--P-GKPLEH--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Heart_kinase> 73 MEILGERTRRHPKQ--VHTMCCTIDEKD--P-GKPLEH--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Lymphocyte_kinase> 58 MEILGERTRRHPKQ--VHTMCCTIDEKD--P-GKPLEH--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 consensus 81 v lQ akkv fn kp dip al ilfv r f lB yl gef kyann g v dt

HeEF-2_kinase> 130 LTPQ--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 MHCK_B> 124 LTPQ--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Melanoma_kinase> 145 LTPQ--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Kidney_kinase> 145 LTPQ--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Muscle_kinase> 154 LTPQ--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Heart_kinase> 150 LTPQ--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 Lymphocyte_kinase> 132 LTPQ--EFTLVQCTNS--EHDLEKSL--SELPEVMTWSSIQ--TVLHCLRE
 consensus 161 afshwtyeyt g llvvdLqg vg d lTdpqi t d g fg gulg gm F H CN C

HeEF-2_kinase> 196 ESEGAPP
 MHCK_B> 190 QYNNQSE
 Melanoma_kinase> 216 EKKCPDR
 Kidney_kinase> 216 EKKCPDR
 Muscle_kinase> 221 ELGCTPR
 Heart_kinase> 217 EKKCPDR
 Lymphocyte_kinase> 206 EKKCPDR
 consensus 241 r l L i

PHYLOGENETIC TREE OF ALPHA-KINASES

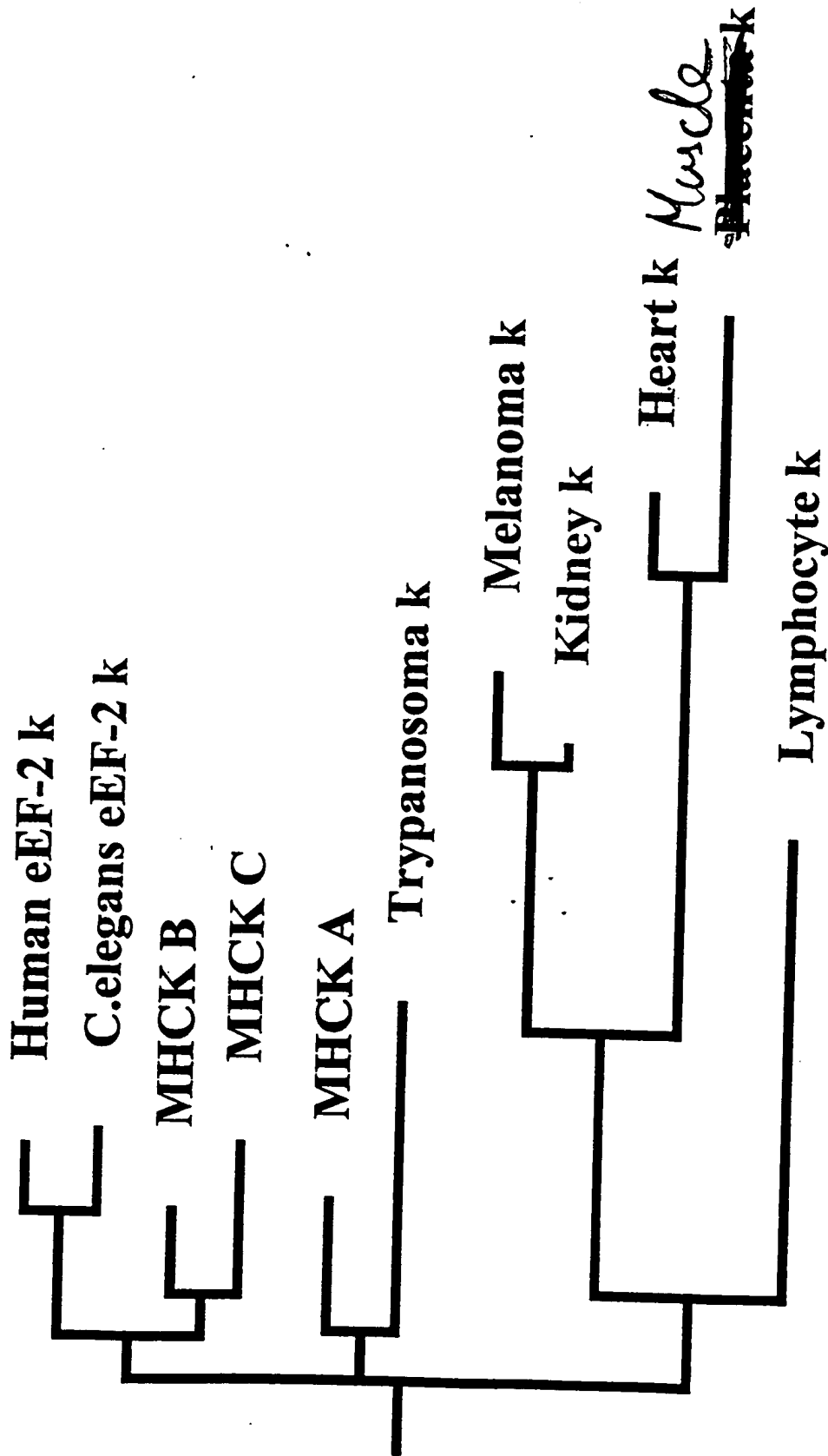
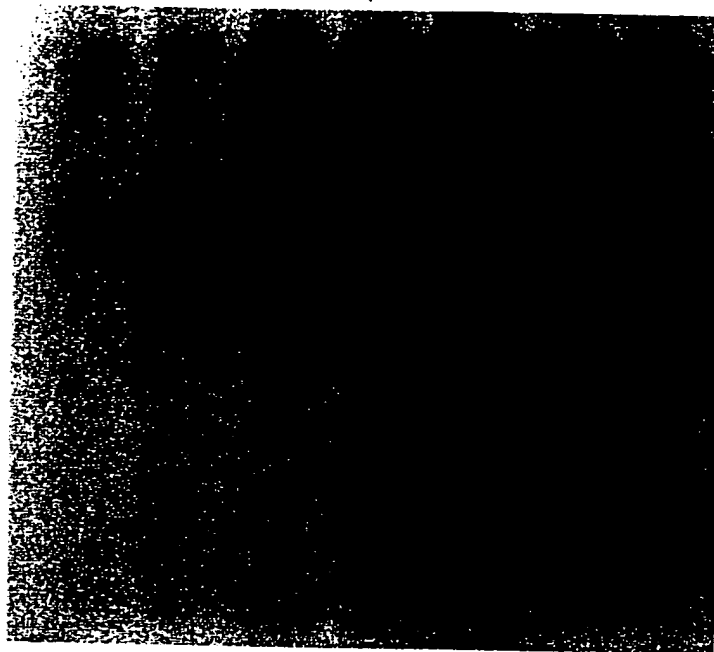


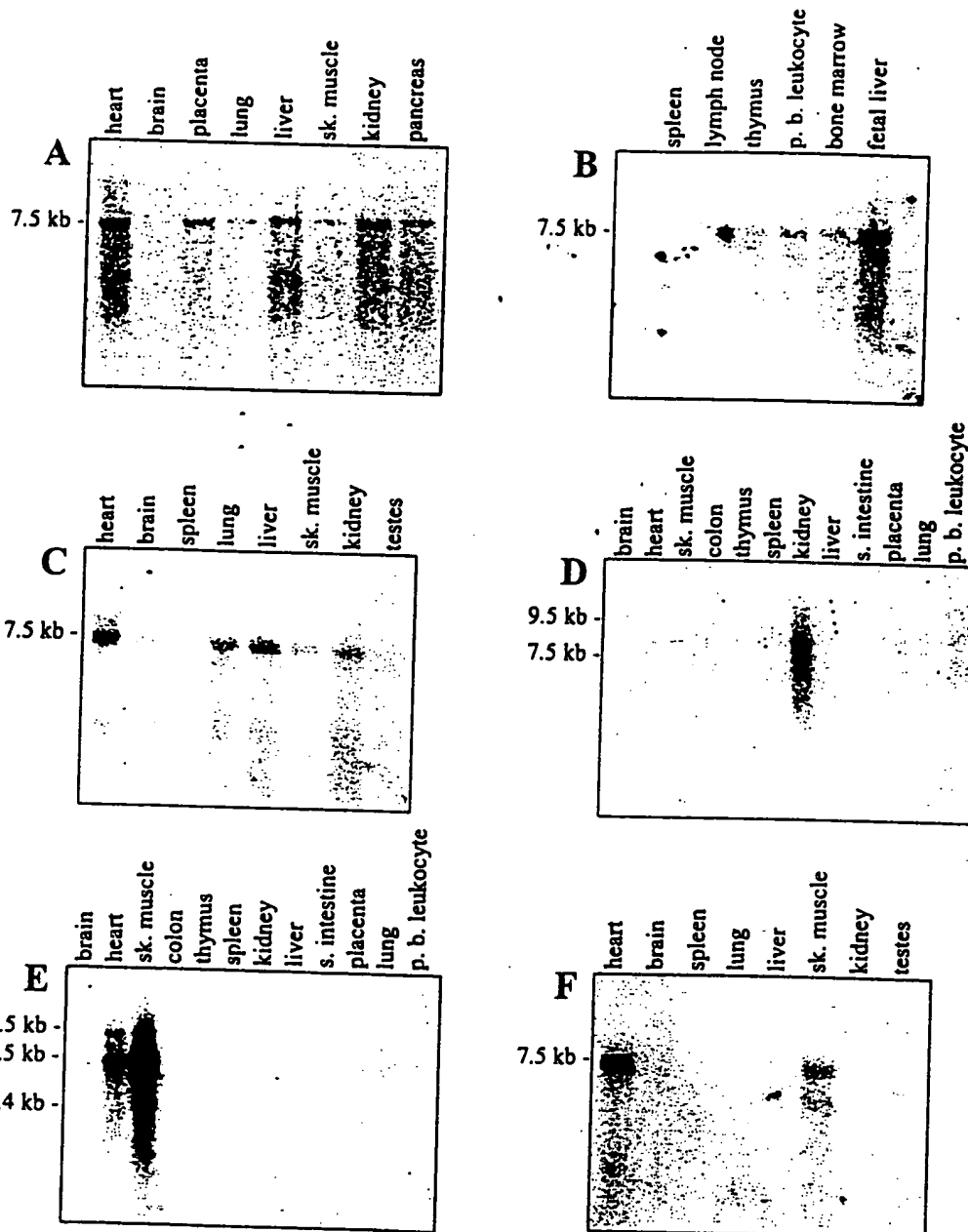
FIGURE 13

Time curve

melanoma kinase
catalytic domain.

0' 5' 10' 20' 30' 60' 120'





NK> NSKNSMTE...
KK> NSKNSMTE...
ME> NSKNSMTE...

NK> 41 KYDOL...
KK> 41 KYDOL...
ME> 41 KYDOL...

NK> 121...
KK> 121...
ME> 121...

NK> 141...
KK> 141...
ME> 141...

NK> 211...
KK> 211...
ME> 211...

NK> 301...
KK> 301...
ME> 301...

NK> 351...
KK> 351...
ME> 351...

NK> 421...
KK> 421...
ME> 421...

NK> 431...
KK> 431...
ME> 431...

NK> 481...
KK> 481...
ME> 481...

NK> 541...
KK> 541...
ME> 541...

NK> 581...
KK> 581...
ME> 581...

NK> 641...
KK> 641...
ME> 641...

NK> 731...
KK> 731...
ME> 731...

NK> 751...
KK> 751...
ME> 751...

NK> 841...
KK> 841...
ME> 841...

NK> 871...
KK> 871...
ME> 871...

NK> 931...
KK> 931...
ME> 931...

NK> 991...
KK> 991...
ME> 991...

NK> 1071...
KK> 1071...
ME> 1071...

NK> 1091...
KK> 1091...
ME> 1091...

NK> 1131...
KK> 1131...
ME> 1131...

NK> 1151...
KK> 1151...
ME> 1151...

NK> 1241...
KK> 1241...
ME> 1241...

NK> 1251...
KK> 1251...
ME> 1251...

NK> 1291...
KK> 1291...
ME> 1291...

NK> 1311...
KK> 1311...
ME> 1311...

NK> 1331...
KK> 1331...
ME> 1331...

NK> 1381...
KK> 1381...
ME> 1381...

NK> 1431...
KK> 1431...
ME> 1431...

NK> 1461...
KK> 1461...
ME> 1461...

NK> 1501...
KK> 1501...
ME> 1501...

NK> 1541...
KK> 1541...
ME> 1541...

NK> 1601...
KK> 1601...
ME> 1601...

NK> 1631...
KK> 1631...
ME> 1631...

NK> 1661...
KK> 1661...
ME> 1661...

NK> 1721...
KK> 1721...
ME> 1721...

NK> 1781...
KK> 1781...
ME> 1781...

NK> 1841...
KK> 1841...
ME> 1841...

NK> 1991...
KK> 1991...
ME> 1991...

Figure 17

601-1-098UP

Figure 17

601-1-098C/P

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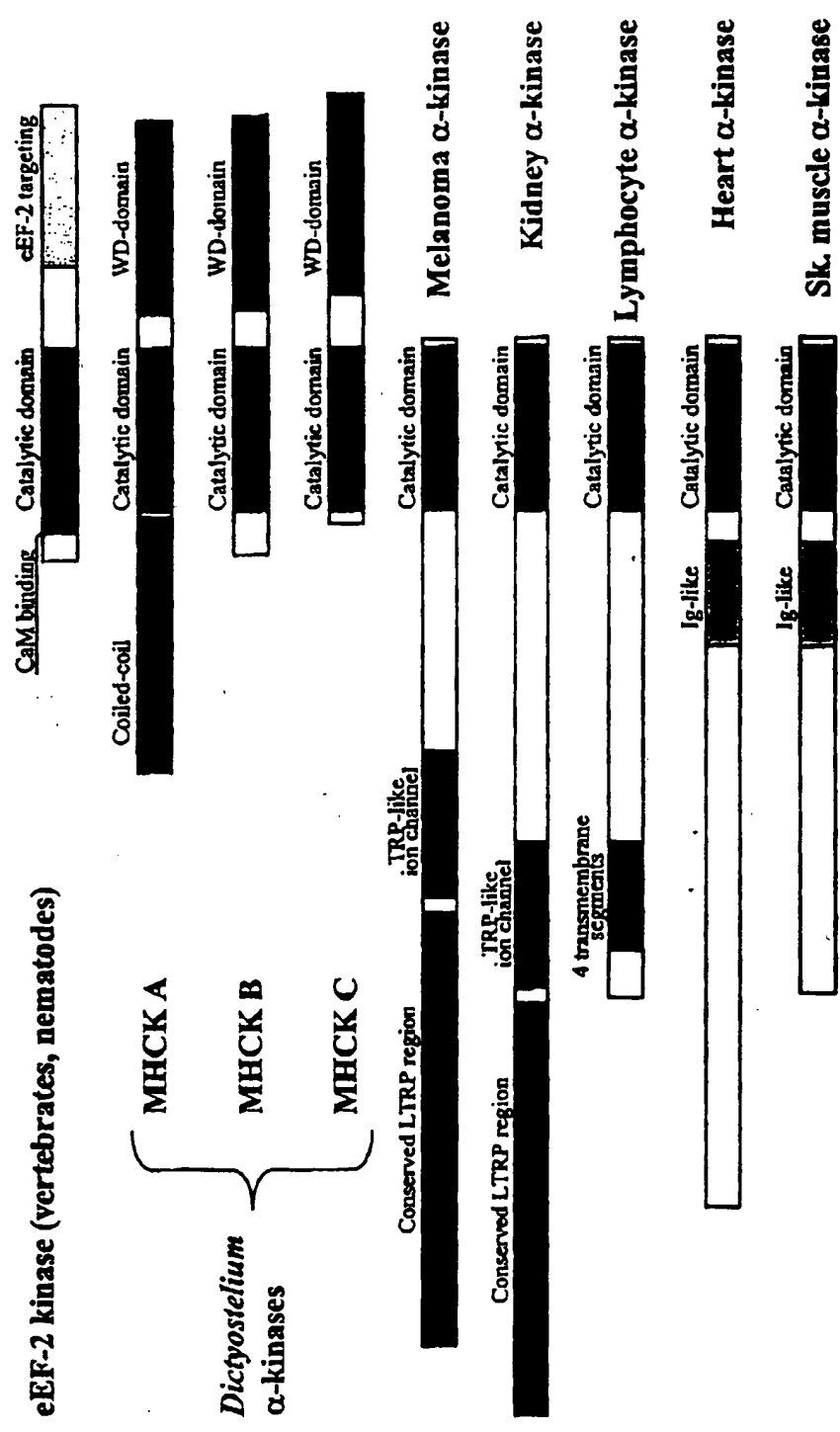


Figure 18

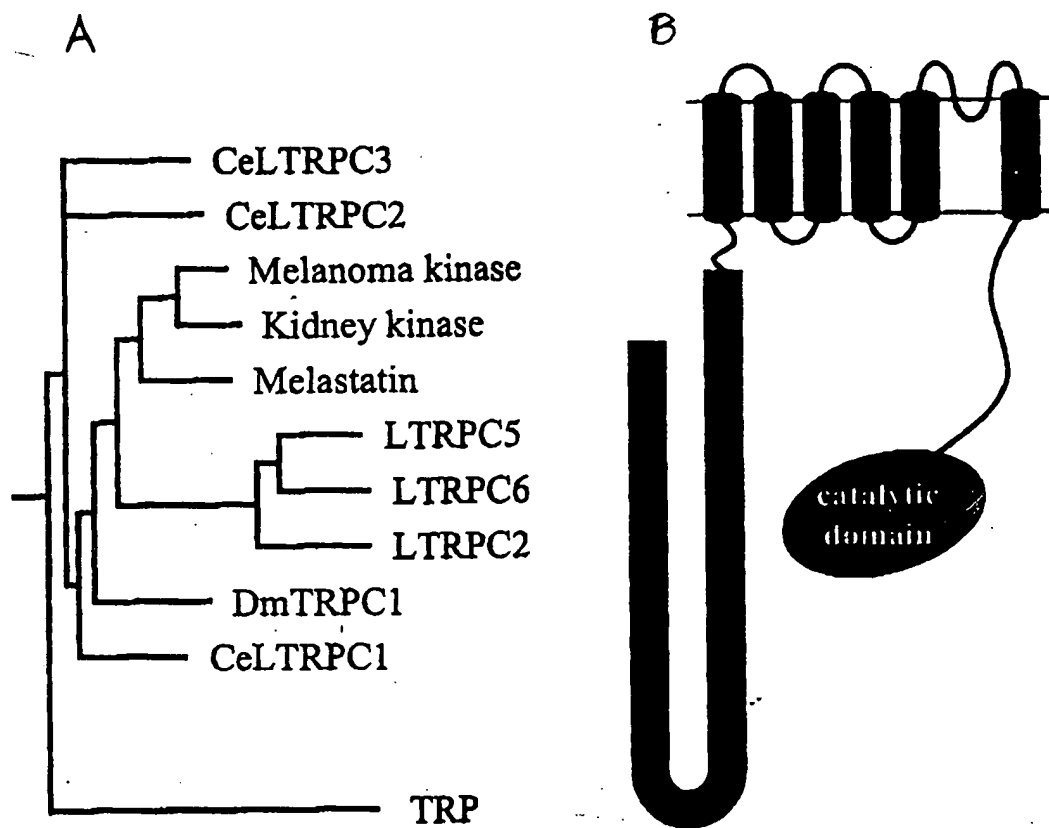


Figure 19A&B